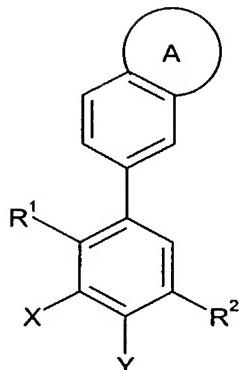


CLAIMS

1. A compound of formula (I):



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(I)

wherein

A is a fused 5-membered heteroaryl ring substituted by $-(CH_2)_m$ aryl or $-(CH_2)_m$ heteroaryl wherein the aryl or heteroaryl is optionally substituted by one or more substituents independently selected from oxo, C₁₋₆alkyl, halogen, -CN, trifluoromethyl, -OR³, -(CH₂)_nCO₂R³, -NR³R⁴, -(CH₂)_nCONR³R⁴, -NHCOR³, -SO₂NR³R⁴, -NHSO₂R³ and -S(O)_pR³, and

10 A is optionally further substituted by one substituent selected from -OR⁵, halogen, trifluoromethyl, -CN, -CO₂R⁵ and C₁₋₆alkyl optionally substituted by hydroxy;

15 R¹ is selected from methyl and chloro;

R² is selected from -NH-CO-R⁶ and -CO-NH-(CH₂)_q-R⁷;

R³ is selected from hydrogen, -(CH₂)_rC₃₋₇cycloalkyl, -(CH₂)_rheterocyclyl, -(CH₂)_raryl, and C₁₋₆alkyl optionally substituted by up to two substituents independently selected from -OR⁸ and -NR⁸R⁹,

20 R⁴ is selected from hydrogen and C₁₋₆alkyl, or

R³ and R⁴, together with the nitrogen atom to which they are bound, form a 5- or 6-membered heterocyclic ring optionally containing one additional heteroatom selected from oxygen, sulfur and N-R¹⁰;

25 R⁵ is selected from hydrogen and C₁₋₆alkyl;

R⁶ is selected from hydrogen, C₁₋₆alkyl, -(CH₂)_q-C₃₋₇cycloalkyl, trifluoromethyl, -(CH₂)_sheteroaryl optionally substituted by R¹¹ and/or R¹², and -(CH₂)_sphenyl optionally substituted by R¹¹ and/or R¹²;

30 R⁷ is selected from hydrogen, C₁₋₆alkyl, C₃₋₇cycloalkyl, -CONHR¹³, phenyl optionally substituted by R¹¹ and/or R¹², and heteroaryl optionally substituted by R¹¹ and/or R¹²;

R⁸ and R⁹ are each independently selected from hydrogen and C₁₋₆alkyl;

R¹⁰ is selected from hydrogen and methyl;

R¹¹ is selected from C₁-6alkyl, C₁-6alkoxy, -(CH₂)_q-C₃-7cycloalkyl, -CONR¹³R¹⁴, -NHCOR¹⁴, halogen, -CN, -(CH₂)_tNR¹⁵R¹⁶, trifluoromethyl, phenyl optionally substituted by one or more R¹² groups, and heteroaryl optionally substituted by one or more R¹² groups;

R¹² is selected from C₁-6alkyl, C₁-6alkoxy, halogen, trifluoromethyl, and -(CH₂)_tNR¹⁵R¹⁶;

R¹³ and R¹⁴ are each independently selected from hydrogen and C₁-6alkyl, or

R¹³ and R¹⁴, together with the nitrogen atom to which they are bound, form a 5- or 6-membered heterocyclic ring optionally containing one additional heteroatom selected from oxygen, sulfur and N-R¹⁰, wherein the ring may be substituted by up to two C₁-6alkyl groups;

R¹⁵ is selected from hydrogen, C₁-6alkyl and -(CH₂)_q-C₃-7cycloalkyl optionally substituted by C₁-6alkyl,

R¹⁶ is selected from hydrogen and C₁-6alkyl, or

R¹⁵ and R¹⁶, together with the nitrogen atom to which they are bound, form a 5- or 6-membered heterocyclic ring optionally containing one additional heteroatom selected from oxygen, sulfur and N-R¹⁰;

X and Y are each independently selected from hydrogen, methyl and halogen;

m, n, p and q are each independently selected from 0, 1 and 2;

r and s are each independently selected from 0 and 1; and

t is selected from 0, 1, 2 and 3;

with the proviso that when A is substituted by -(CH₂)_mheteroaryl and m is 0, the -(CH₂)_mheteroaryl group is not a 5-membered heteroaryl ring optionally substituted by C₁-alkyl;

or a pharmaceutically acceptable derivative thereof.

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2. A compound according to claim 1 wherein A is a fused 5-membered heteroaryl ring containing up to two heteroatoms independently selected from oxygen and nitrogen.

30

3. A compound according to claim 1 or claim 2 wherein R¹ is methyl.

35

4. A compound according to any one of the preceding claims wherein R² is -CO-NH-(CH₂)_q-R⁷.

40

5. A compound according to any one of the preceding claims wherein A is substituted by -(CH₂)_mheteroaryl wherein the heteroaryl is a 5- or 6-membered heteroaryl ring containing up to two heteroatoms independently selected from oxygen and nitrogen.

6. A compound according to claim 5 wherein the heteroaryl is optionally substituted by one or two substituents independently selected from oxo, C₁-6alkyl, halogen, -OR³, -NR³R⁴ and -(CH₂)_nCONR³R⁴.

7. A compound according to claim 6 wherein the heteroaryl is substituted by one or two substituents independently selected from oxo and C₁₋₆alkyl.

8. A compound according to any one of claims 1 to 4 wherein A is substituted by -
5 (CH₂)_maryl wherein the aryl is phenyl.

9. A compound according to claim 8 wherein the aryl is substituted by one or two substituents independently selected from C₁₋₆alkyl, halogen, -CN, trifluoromethyl, -OR³, -NR³R⁴, -(CH₂)_nCONR³R⁴ and -S(O)_pR³.

10

10. A compound according to any one of the preceding claims wherein X is hydrogen or fluorine.

15

11. A compound according to claim 1 substantially as hereinbefore defined with reference to any one of Examples 1 to 82, or a pharmaceutically acceptable derivative thereof.

12. A compound selected from:

N-cyclopropyl-3-fluoro-4-methyl-5-(1-phenyl-1H-indazol-5-yl)benzamide;

N-cyclopropyl-3-fluoro-5-[1-(4-fluorophenyl)-1H-indazol-5-yl]-4-methylbenzamide;

20

N-cyclopropyl-3-fluoro-5-[1-(4-fluoro-2-methylphenyl)-1H-indazol-5-yl]-4-methylbenzamide;

N-cyclopropyl-3-fluoro-4-methyl-5-{1-[4-(4-morpholinyl)phenyl]-1H-indazol-5-yl}benzamide;

N-ethyl-3-fluoro-4-methyl-5-(1-phenyl-1H-indazol-5-yl)benzamide;

N-(cyclopropylmethyl)-3-fluoro-4-methyl-5-(1-phenyl-1H-indazol-5-yl)benzamide;

N-cyclopropyl-3-fluoro-4-methyl-5-{1-[4-(methylsulfonyl)phenyl]-1H-indazol-5-yl}benzamide;

25

N-cyclopropyl-3-fluoro-4-methyl-5-(1-{4-[2-(methylamino)-2-oxoethyl]phenyl}-1H-indazol-5-yl)benzamide;

N-cyclopropyl-3-[1-(4-{[2-(dimethylamino)ethyl]amino}phenyl)-1H-indazol-5-yl]-5-fluoro-4-methylbenzamide;

30

N-cyclopropyl-3-fluoro-4-methyl-5-{1-[4-(tetrahydro-2H-pyran-4-ylamino)phenyl]-1H-indazol-5-yl}benzamide;

N-cyclopropyl-3-(1-{4-[(2,3-dihydroxypropyl)amino]phenyl}-1H-indazol-5-yl)-5-fluoro-4-methylbenzamide;

35

N-cyclopropyl-3-fluoro-4-methyl-5-{3-[4-(methyloxy)phenyl]-1,2-benzisoxazol-6-yl}benzamide;

N-cyclopropyl-3-fluoro-5-[3-(4-hydroxyphenyl)-1,2-benzisoxazol-6-yl]-4-methylbenzamide;

N-cyclopropyl-3-fluoro-4-methyl-5-{1-[(1-oxido-2-pyridinyl)methyl]-1H-indazol-5-yl}benzamide;

40

N-ethyl-3-[3-(4-fluorophenyl)-1H-indazol-6-yl]-4-methylbenzamide;

N-cyclopropyl-3-[3-(4-fluorophenyl)-1H-indazol-6-yl]-4-methylbenzamide;

N-ethyl-4-methyl-3-{3-[4-(methyloxy)phenyl]-1H-indazol-6-yl}benzamide;

N-cyclopropyl-4-methyl-3-[3-[4-(methyloxy)phenyl]-1*H*-indazol-6-yl]benzamide;
N-(1-ethyl-1*H*-pyrazol-5-yl)-3-fluoro-5-[3-(4-fluorophenyl)-1*H*-indazol-6-yl]-4-methylbenzamide;
3-fluoro-5-[3-(4-fluorophenyl)-1*H*-indazol-6-yl]-4-methyl-N-(1-methyl-1*H*-pyrazol-5-yl)benzamide;
5
N-ethyl-3-fluoro-5-[3-[4-fluoro-2-(methyloxy)phenyl]-1*H*-indazol-6-yl]-4-methylbenzamide;
N-(1,4-dimethyl-1*H*-pyrazol-5-yl)-3-fluoro-5-[3-(4-fluorophenyl)-1*H*-indazol-6-yl]-4-methylbenzamide; and
N-(1,4-dimethyl-1*H*-pyrazol-5-yl)-3-[3-(4-fluorophenyl)-1*H*-indazol-6-yl]-4-methylbenzamide;
10 or a pharmaceutically acceptable derivative thereof.

13. A pharmaceutical composition comprising at least one compound as claimed in any one of claims 1 to 12, or a pharmaceutically acceptable derivative thereof, in association with one or more pharmaceutically acceptable excipients, diluents and/or carriers.

15

14. A compound according to any one of claims 1 to 12, or a pharmaceutically acceptable derivative thereof, for use in therapy.

20

15. A compound as claimed in any one of claims 1 to 12, or a pharmaceutically acceptable derivative thereof, for use in the treatment or prophylaxis of a condition or disease state mediated by p38 kinase activity or mediated by cytokines produced by the activity of p38 kinase.

25

16. A method for treating a condition or disease state mediated by p38 kinase activity or mediated by cytokines produced by the activity of p38 kinase comprising administering to a patient in need thereof a compound as claimed in any one of claims 1 to 12, or a pharmaceutically acceptable derivative thereof.

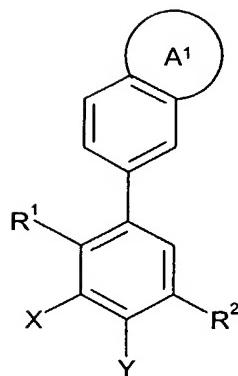
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17. Use of a compound as claimed in any one of claims 1 to 12, or a pharmaceutically acceptable derivative thereof, in the manufacture of a medicament for use in the treatment of a condition or disease state mediated by p38 kinase activity or mediated by cytokines produced by the activity of p38 kinase.

35

18. A process for preparing a compound of formula (I) as claimed in any one of claims 1 to 12, or a pharmaceutically acceptable derivative thereof, which comprises

(a) reacting a compound of formula (II)



(II)

in which R^1 , R^2 , X and Y are as defined in claim 1 and A^1 is an unsubstituted fused 5-membered heteroaryl ring with a halide derivative of formula (IIIA) or (IIIB)

5

 $\text{Z-(CH}_2\text{)}_m\text{aryl}$

(IIIA)

 $\text{Z-(CH}_2\text{)}_m\text{heteroaryl}$

(IIIB)

10

in which $-(\text{CH}_2)_m\text{aryl}$ and $-(\text{CH}_2)_m\text{heteroaryl}$ are as defined in claim 1 and Z is halogen, in the presence of a base,

or, when A is substituted by $-(\text{CH}_2)_m\text{aryl}$ wherein m is 0, reacting the compound of formula (II) with a boronic acid compound of formula (IV)

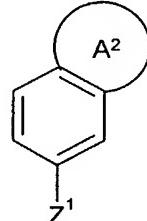
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 $(\text{HO})_2\text{B-(CH}_2\text{)}_m\text{aryl}$

(IV)

in which $-(\text{CH}_2)_m\text{aryl}$ is as defined in claim 1,

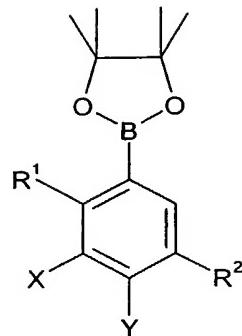
20 (b) reacting a compound of formula (V)



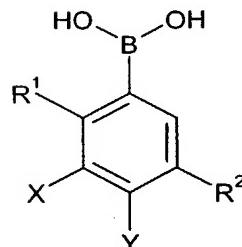
(V)

in which A^2 is A as defined in claim 1 and Z^1 is halogen, with a compound of formula (VI A) or (VI B)

25



(VIA)

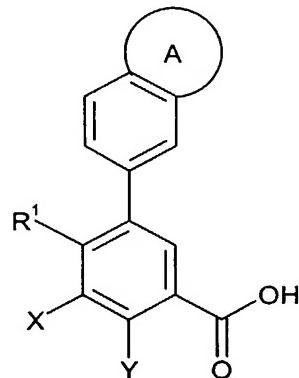


(VIB)

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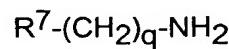
in which R¹, R², X and Y are as defined in claim 1,
in the presence of a catalyst;

- 10 (c) reacting a compound of formula (XVI)



(XVI)

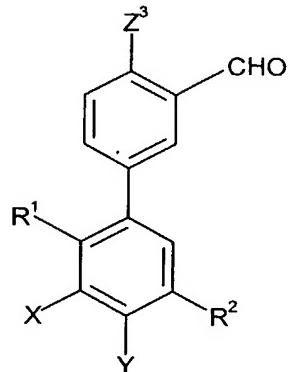
- 15 in which A, R¹, X and Y are as defined in claim 1,
with an amine compound of formula (XV)



(XV)

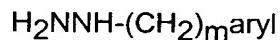
- 20 in which R⁷ and q are as defined in claim 1,
under amide forming conditions;

d) when A is a fused pyrazolyl, reacting a compound of formula (XVII)

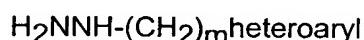


5 (XVII)

in which R¹, R², X and Y are as defined in claim 1 and Z³ is halogen, with a hydrazine derivative of formula (VIIIA) or (VIIIB)



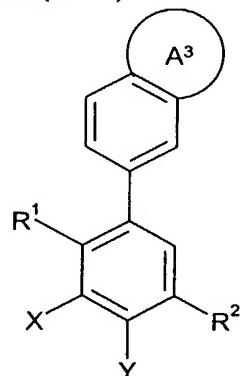
10 (VIIIA)



(VIIIB)

15 in which -(CH₂)_maryl and -(CH₂)_mheteroaryl are as defined in claim 1;

(e) reacting a compound of formula (XVIII)



(XVIII)

20 in which R¹, R², X and Y are as defined in claim 1 and A³ is a fused 5-membered heteroaryl ring substituted by halogen, with a suitable boronic acid derivative; or

(f) final stage modification of one compound of formula (I) as defined in claim 1 to give another compound of formula (I) as defined in claim 1.